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What Is Claimed Is:

1. A compound of Formula I:

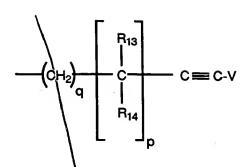
wherein A is selected from methylene, CO, SO and SO₂; wherein X is selected from oxygen atom, methylene and NR₁₀ with R₁₀ selected from hydrido, alkyl and benzyl; wherein each of R₁ and R₉ is a group independently selected from hydrido, alkyl, cycloalkyl, alkoxyacyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl,

loweralkanoyl, haloalkylacyl, phenyl, benzyl, naphthyl, and naphthylmethyl, any one of which groups having a substitutable position may be optionally substituted with one or more radicals selected from alkyl, alkoxy, alkenyl, alkynyl, halo, haloalkyl, cyano and phenyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an N-oxide; wherein R2 is

selected from hydrido, alkyl, dialkylaminoalkyl, alkylacylaminoalkyl, benzyl and cycloalkyl; wherein R3 is selected from alkyl, cycloalkylalkyl, acylaminoalkyl, phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and heterocycliccycloalkyl, wherein the cyclic portion of any of said phenylalkyl, naphthylmethyl, aryl,

of said phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and heterocycliccycloalkyl groups may be substituted by one or more radicals selected from halo, hydroxy, alkoxy and alkyl; wherein each of R4 and R6 is independently selected from hydrido, alkyl, benzyl and cycloalkyl; wherein each of R5 and R8 is independently

selected from



wherein V is selected from hydrido, alkyl, cycloalkyl, haloalkyl, benzyl and phenyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, phenyl, heterocyclic, heterocyclicalkyl and heterocycliccycloalkyl; wherein R7 is selected from substituted or unsubstituted alkyl, cycloalkyl, phenyl, cycloalkylalkyl and phenylalkyl, any one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo, haloalkyl, alkenyl, alkynyl and cyano; wherein each of R_{11} and R_{12} independently selected from hydrido, alkyl, haloalkyl, dialkylamino and phenyl; and wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

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2. Compound of Claim I wherein A is selected from methylene, CO, SO and SO2; wherein X is selected from oxygen atom, methylene and NR₁₀ with R₁₀ selected from hydrido, alkyl and benzyl; wherein each of R₁ and R₉ is independently selected from hydrido, lower alkyl, haloalkyl, cycloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, alkoxyacyl, phenyl and benzyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein each of R₂, R₄ and R₆ is independently selected from hydrido and alkyl; wherein R₃ is selected from phenylalkyl, naphthylmethyl, cyclohexylalkyl, cyclopentylalkyl,

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15 2 heteroarylalkyl and heteroarylcycloalkyl; wherein each of R5 and R8 is independently selected from

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wherein V is selected from hydrido, alkyl, haloalkyl, benzyl and phenyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, heteroaryl, heteroarylalkyl and heteroarylcycloalkyl; wherein R7 is selected from substituted or unsubstituted cyclohexylmethyl and benzyl, either one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo and haloalkyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

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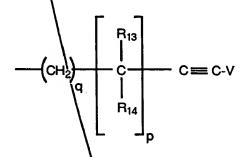
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from methylene, CO, SO and SO2; wherein X is selected from oxygen atom, methylene and NR10 with R10 selected from hyrido, alkyl and benzyl; wherein each of R1 and R9 is independently selected from hydrido, alkyl, alkoxyacyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl and benzyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an N-oxide; wherein each of R2, R4 and R6 is independently selected from hydrido and alkyl; wherein R3 is selected from benzyl, phenethyl, cyclohexylmethyl, phenpropyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl,

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piperidinylmethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, furanylmethyl, furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazolemethyl, pyridazinemethyl, pyridazinemethyl, pyridazinemethyl, pyrazinemethyl and pyrazineethyl; wherein each of R5 and R8 is independently selected from

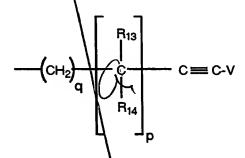


wherein V is selected from hydrido, alkyl and haloalkyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, thiazole and thiazolemethyl; wherein R7 is cyclohexylmethyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

from CO and SO₂; wherein X is selected from oxygen atom, methylene and NR₁₀ with R₁₀ selected from hydrido and methyl; wherein each of R₁ and R₉ is independently selected from hydrido, lower alkyl, alkoxyacyl, alkoxycarbonyl, benzyloxycarbonyl, haloalkyl and benzyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein R₂ is selected from hydrido, methyl, ethyl and isopropyl; wherein R₃ is selected from benzyl, phenethyl,

cyclohexylmethyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl, pyrazolemethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazolemethyl, imidazolemethyl, imidazolemethyl, imidazolemethyl, thienylmethyl, thienylmethyl, furanylmethyl, oxazolemethyl, oxazolemethyl, isoxazolemethyl, pyridazinemethyl, pyridazinemethyl, pyridazinemethyl, pyridazinemethyl;

wherein each of R4 and R6 is independently selected from hydrido and methyl; wherein each of R5 and R8 is independently selected from



wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl and alkynyl; wherein R7 is cyclohexylmethyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

5. Compound of Claim 4 wherein A is selected from CO and SO2; wherein X is selected from oxygen atom and methylene; wherein each of R1 and R9 is independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen

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to form an N-oxide; wherein R2 is selected from hydrido, methyl, ethyl and isopropyl; wherein R3 is selected from benzyl, cyclohexylmethyl, phenethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazolemethyl, imidazolemethyl, imidazolemethyl, thienylmethyl, thienylethyl, furanylethyl, thienylethyl, oxazolemethyl, isoxazolemethyl, oxazolemethyl, pyridazinemethyl, pyridazinemethyl, pyridazinemethyl, pyridazinemethyl, wherein each of R5 and R8 is independently selected from

 $-(CH_2) = \begin{pmatrix} R_{13} \\ C \\ R_{14} \end{pmatrix} = C = C-V$

wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R13 and R14 is a radical independently selected from hydrido, methyl, ethyl, propyl and ethynyl; wherein R7 is cyclohexylmethyl; wherein each of R4 and R6 is independently selected from hydrido and methyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein m is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically acceptable salt thereof.

6. Compound of Claim 5 wherein A is selected from CO and SO2; wherein X is selected from oxygen atom and methylene; wherein each of R1 and R9 is a group independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein

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the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an N-oxide; wherein R2 is selected from hydrido, methyl, ethyl and isopropyl; wherein R3 is selected from benzyl, cyclohexylmethyl, phenethyl, imidazolemethyl, pyridylmethyl and 2-pyridylethyl; wherein each of R5 and R8 is independently selected from

wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R13 and R14 is a radical independently selected from hydrido, methyl and ethynyl; wherein R7 is cyclohexylmethyl; wherein each of R4 and R6 is independently selected from hydrido and methyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl and phenyl; wherein m is zero; wherein n is a number selected from zero through three; wherein p is a number selected from one through three; and wherein q is zero or one; or a pharmaceutically-acceptable salt thereof.

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7. Compound of Claim 6 selected from compounds, their tautomers, and the pharmaceutically-acceptable esters and salts thereof, of the group consisting of

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8. Compound of Claim 6 which is N1-[1R*-[[[1s,1R*-(cyclohexylmethyl)-2s*,3R*-dihydroxy-hexynyl]amino]carbonyl]-3-butynyl]-N4-[2-(dimethylamino)ethyl]-N4-methyl-2s*-(phenylmethyl)butanediamide or a pharmaceutically-acceptable salt thereof.

9. Compound of Claim 6 which is [1R*-[[[1R*-[[[1S,1R*-(cyclohexylmethyl)-2S*,3R*-dihydroxyhexynyl]amino]carbonyl]-3-butynyl]amino]carbonyl]-2phenylethyl)[2-(dimethylamino)ethyl]methylcarbamate or a pharmaceutically-acceptable salt thereof.

CF3 N Compound of Claim 6 which is

11. Compound of Claim 6\which is

or a pharmaceutically-acceptable salt the reof.

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A pharmaceutical composition comprising 12. a therapeutically-effective amount of a renin-inhibiting compound and a pharmaceutically-acceptable carrier or diluent, said renin-inhibiting compound selected from a family of compounds of Formula I:

wherein A is selected from methylene, CO, SO and SO2; wherein X is selected from paygen atom, methylene and \mathtt{NR}_{10} with \mathtt{R}_{10} selected from hydrido, alkyl and benzyl; wherein each of R_1 and R_9 is a group independently selected from hydrido, alkyl, cycloalkyl, alkoxyacyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, haloalkylacyl, phenyl, benzyl, naphthyl, and naphthylmethyl, any one of which groups having a substitutable position may be optionally substituted with one or more radicals selected from alkyl, alkoxy, alkenyl, alkynyl, halo, haloalkyl, cyano and phenyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an N-oxide; wherein R2 is selected from hydrido, alkyl, dialkylaminoalkyl, alkylacylaminoalkyl, benzyl and cycloalkyl; wherein R3 is selected from alkyl, cycloalkylalkyl, acylaminoalkyl, phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and heterocycliccycloalkyl, wherein the cyclic portion of any of said phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and heterocycliccycloalkyl groups may be substituted by one or more radicals selected from halo, hydroxy, alkoxy and alkyl; wherein each of R4 and R6 is independently selected from hydrido, alkyl, benzyl and cycloalkyl; wherein each of R5 and R8 is independently selected from

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wherein V is selected from hydrido, alkyl, cycloalkyl, haloalkyl, benzyl and phenyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, phenyl, heterocyclic, heterocyclicalkyl and heterocycliccycloalkyl; wherein R7 is selected from substituted or unsubstituted alkyl, cycloalkyl, phenyl, cycloalkylalkyl and phenylalkyl, any one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo, haloalkyl, alkenyl, alkynyl and cyano; wherein each of R11 and R12 is independently selected from hydrido, alkyl, haloalkyl, dialkylamino and phenyl; and wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is

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13. The composition of Claim 12 wherein A is selected from methylene, CO, SO and SO2; wherein X is selected from oxygen atom, methylene and NR₁₀ with R₁₀ selected from hydrido, alkyl and benzyl; wherein each of R₁ and R₉ is independently selected from hydrido, lower alkyl, haloalkyl, cycloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, alkoxyacyl, phenyl and

a number selected from zero through five; or a

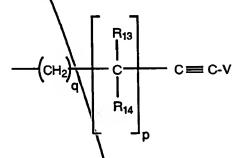
pharmaceutically-acceptable salt thereof.

benzyloxycarbonyl, loweralkanoyl, alkoxyacyl, phenyl and benzyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an

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N-oxide; wherein each of R2, R4 and R6 is independently selected from hydrido and alkyl; wherein R3 is selected from phenylalkyl, maphthylmethyl, cyclohexylalkyl, cyclopentylalkyl, heteroarylalkyl and heteroarylcycloalkyl; wherein each of R5 and R8 is independently selected from



wherein V is selected from hydrido, alkyl, haloalkyl, benzyl and phenyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, heteroaryl, heteroarylalkyl and heteroarylcycloalkyl; wherein R7 is selected from substituted or unsubstituted cyclohexylmethyl and benzyl, either one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo and haloalkyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein m is a number selected from zero through five; wherein m is a number selected from zero through five; and wherein m is a number selected from zero through five; and wherein m is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

14. The composition of Claim 13 wherein A is selected from methylene, CO, SO and SO2; wherein X is selected from oxygen atom, methylene and NR₁₀ with R₁₀ selected from hyrido, alkyl and benzyl; wherein each of R₁ and R₉ is independently selected from hydrido, alkyl, alkoxyacyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl, and benzyl, and wherein the nitrogen

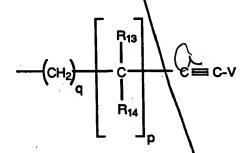
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atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein each of R₂, R₄ and R₆ is independently selected from hydrido and alkyl; wherein R₃ is selected from benzyl, phenethyl, cyclohexylmethyl, phenpropyl, pyrrolidinyl, piperidinyl,

cyclohexylmethyl, phenpropyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl, pyrazolemethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazolemethyl, imidazolemethyl, imidazolemethyl, thienylmethyl, thienylethyl, furanylmethyl, furanylethyl, oxazolemethyl, oxazolemethyl, isoxazolemethyl, isoxazolemethyl, pyridazinemethyl, pyridazinemethyl, pyridazinemethyl, pyrazinemethyl and pyrazineethyl; wherein each of R5 and R8 is independently selected from



wherein V is selected from hydrido, alkyl and haloalkyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, thiazole and thiazolemethyl; wherein R₇ is cyclohexylmethyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

15. The composition of Claim 14 wherein A is selected from CO and SO2; wherein X is selected from oxygen atom, methylene and NR₁₀ with R₁₀ selected from

hydrido and methyl; wherein each of R1 and R9 is independently selected from hydrido, lower alkyl, alkoxyacyl, alkoxydarbonyl, benzyloxycarbonyl, haloalkyl and benzyl, and wherein the nitrogen atom to which R1 and Rg are attached may be combined with oxygen to form an N-oxide; wherein Ro is selected from hydrido, methyl, ethyl and isopropyl; wherein R3 is selected from benzyl, phenethyl, cyclohexylmethyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, thienylethyl, furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of R4 and R6 is independently selected from hydrido and methyl; wherein each of R5 and R8 is independently selected from

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wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl and alkynyl; wherein R7 is cyclohexylmethyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

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The composition of Claim 15 wherein A is selected from CO and SO2; wherein X is selected from oxygen atom and methylene; wherein each of R1 and R9 is independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an N-oxide; wherein R2 is selected from hydrido, methyl, ethyl and isopropyl; wherein R3 is selected from benzyl, cyclohexylmethyl, phenethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, thienylethyl, furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, \isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of Rt and Rg is independently selected from

$$\begin{array}{c|c}
\hline
 & CH_2 & \hline
 & CH_$$

wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R13 and R14 is a radical independently selected from hydrido, methyl, ethyl, propyl and ethynyl; wherein R7 is cyclohexylmethyl; wherein each of R4 and R6 is independently selected from hydrido and methyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through

five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

17. The composition of Claim 16 wherein A is selected from CO and SO2; wherein X is selected from oxygen atom and methylene; wherein each of R1 and R9 is a group independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an N-oxide; wherein R2 is selected from hydrido, methyl, ethyl and isopropyl; wherein R3 is selected from benzyl, cyclohexylmethyl, phenethyl, imidazolemethyl, pyridylmethyl and 2-pyridylethyl; wherein each of R5 and R8 is independently selected from

$$\begin{array}{c|c}
\hline
 & CH_2 & \hline
 & CH_$$

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wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R13 and R14 is a radical independently selected from hydrido, methyl and ethynyl; wherein R7 is cyclohexylmethyl; wherein each of R4 and R6 is independently selected from hydrido and methyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl and phenyl; wherein m is zero; wherein n is a number selected from zero through three; wherein p is a number selected from one through three; and wherein q is zero or one; or a pharmaceutically-acceptable salt thereof.

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18. The composition of Claim 17 wherein said renin-inhibiting compound is selected from compounds, their tautomers, and the pharmaceutically-acceptable esters and salts thereof of the group consisting of

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C≡CH H OH N ▼ Ē Он ·C≡CH H QH c≡ch h QH and

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19. The composition of Claim 17 wherein said renin-inhibiting compound is Nl-[1R*-[[[1S,1R*-(cyclohexylmethyl)-2S*,3R*-dihydroxy-hexynyl]amino]carbonyl]-3-butynyl]-N4-[2-(dimethylamino)ethyl]-N4-methyl-2S*-(phenylmethyl)butanediamide or a pharmaceutically-acceptable salt thereof.

20. The composition of Claim 17 wherein said renin-inhibiting compound is [lR*-[[lR*-[[lS,lR*-(cyclohexylmethyl)-25*,3R*-dihydroxy-hexynyll]amino]carbonyl]-3-butynyl]amino]carbonyl]-2-phenylethyl)[2-(dimethylamino)ethyl]methylcarbamate or a pharmaceutically-acceptable salt thereof.

21. The composition of Claim 17 wherein said renin-inhibiting compound is

22. The composition of Claim 17 wherein said renin-inhibiting compound is

or a pharmaceutically-acceptable salt thereof.

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23. A therapeutic method for treating a circulatory disorder or a circulatory-related disorder, said method comprising administering to a subject susceptible to or afflicted with such disorder a therapeutically-effective amount of an active compound

of Formula I:

wherein A is selected from methylene, CO, SO and SO2: wherein X is selected from oxygen atom, methylene and NR_{10} with R_{10} selected from hydrido, alkyl and benzyl; wherein each of R₁ and R₉ is-a group independently selected from hydrido, alkyl, eycloalkyl, alkoxyacyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, haloalkylacyl, phenyl, benzyl, naphthyl, and naphthylmethyl, any one of which groups having a substitutable position may be optionally substituted with one or more radicals selected from alkyl, alkoxy, alkenyl, alkynyl, halo, haloalkyl, cyano and phenyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an N-oxide; wherein R2 is selected from hydrido, alkyl, dialkylaminoalkyl, alkylacylaminoalkyl, benzyl and cycloalkyl; wherein R3 is selected from alkyl, cycloalkylalkyl, acylaminoalkyl, phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and heterocycliccycloalkyl, wherein the cyclic portion of any of said phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and heterocycliccycloalkyl groups may be substituted by one or more radicals selected from halo, hydroxy, alkoxy and alkyl; wherein each of R4 and R6 is independently selected from hydrido\ alkyl, benzyl and cycloalkyl; wherein each of R5 and R8 is independently selected from

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TJ CD wherein V is selected from hydrido, alkyl, cycloalkyl, haloalkyl, benzyl and phenyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, phenyl, heterocyclic, heterocyclicalkyl and heterocycliccycloalkyl; wherein R7 is selected from substituted or unsubstituted alkyl, cycloalkyl, phenyl, cycloalkylalkyl and phenylalkyl, any one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo, haloalkyl, alkenyl, alkynyl and cyano; wherein each of R11 and R12 is independently selected from hydrido, alkyl, \haloalkyl, dialkylamino and phenyl; and wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a

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24. The method of Claim 23 wherein A is selected from methylene, CO, SO and SO2; wherein X is selected from oxygen atom, methylene and NR₁₀ with R₁₀ selected from hydrido, alkyl and benzyl; wherein each of R₁ and R₉ is independently selected from hydrido, lower alkyl, haloalkyl, cycloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, alkoxyacyl, phenyl and

pharmaceutically-acceptable salt thereof.

benzyloxycarbonyl, loweralkanoyl, alkoxyacyl, phenyl and benzyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an

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N-oxide; wherein each of R2, R4 and R6 is independently selected from hydrido and alkyl; wherein R3 is selected from phenylalkyl, naphthylmethyl, cyclohexylalkyl, cyclopentylalkyl, heteroarylalkyl and heteroarylcycloalkyl; wherein each of R5 and R8 is independently selected from

$$-(CH_2)_{q} = C = C-V$$

wherein V is selected from hydrido, alkyl, haloalkyl, benzyl and phenyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, heteroaryl, heteroarylalkyl and heteroarylcycloalkyl; wherein R7 is selected from substituted or unsubstituted cyclohexylmethyl and benzyl, either one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo and haloalkyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

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The method of Claim 24 wherein A is selected from mathylene, CO, SO and SO2; wherein X is selected from oxygen atom, methylene and NR10 with R10 selected from hyrido, alkyl and benzyl; wherein each of R1 and R9 is independently selected from hydrido, alkyl, 5 alkoxyacyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl and benzyl, and wherein the nitrogen atom to which R1 and Rg are attached may be combined with oxygen to form an N-oxide; wherein each of R2, R4 and R6 is independently selected from hydrido and alkyl; wherein 10 R3 is selected from benzyl, phenethyl, cyclohexylmethyl, phenpropyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, thienylethyl, furanylmethyl, furanylethyl (oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of R5 and R8 is independently selected from

$$-\left(CH_{2}\right)_{q} \begin{bmatrix} R_{13} \\ C \\ R_{14} \end{bmatrix}_{p} C = C-V$$

wherein V is selected from hydrido, \alkyl and haloalkyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl, alkenyl alkynyl, thiazole and thiazolemethyl; wherein R7 is cyclohexylmethyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero

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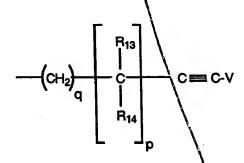
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The method of Claim 25 wherein A is selected from CO and SO2; wherein X is selected from oxygen atom, methylene and $\sum NR_{10}$ with R_{10} selected from hydrido and methyl; wherein each of R1 and R9 is independently selected from hydrido, lower alkyl, alkoxyacyl, alkoxycarbonyl, benzyloxycarbonyl, haloalkyl and benzyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an N-oxide; wherein R2 is selected from hydrido, methyl, ethyl and isopropyl; wherein R3 is selected from benzyl, phenethyl, cyclohexylmethyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, thienylethyl, furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of R4 and R6 is independently selected from hydrido and methyl; wherein each of R5 and R8 is independently selected from



wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl and alkynyl; wherein R7 is cyclohexylmethyl; wherein each of

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R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

27. The method of Claim 26 wherein A is selected from CO and \$02; wherein X is selected from oxygen atom and methylene; wherein each of R1 and R9 is independently selected \from hydrido, methyl, ethyl, n-propyl, isopropyl, behzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an N-oxide; wherein R2 is selected from hydrido, methyl, ethyl and isopropyl; wherein R3 is selected from benzyl, cyclohexylmethyl, phenethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, thienylethyl, furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of \$\,\frac{1}{2}\$ and R8 is independently selected from

$$-\left(CH_{2}\right)_{q} \begin{bmatrix} R_{13} \\ C \\ R_{14} \end{bmatrix}_{p} C = C-V$$

wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, methyl, ethyl, propyl and ethynyl; wherein R₇ is

cyclohexylmethyl; wherein each of R4 and R6 is independently selected from hydrido and methyl; wherein each of R11 and R12 is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

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28. The method of Claim 27 wherein A is selected from CO and SO2; wherein X is selected from oxygen atom and methylene; wherein each of R1 and R9 is a group independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R1 and R9 are attached may be combined with oxygen to form an N-oxide; wherein R2 is selected from hydrido, methyl, ethyl and isopropyl; wherein R3 is selected from benzyl, cyclohexylmethyl, phenethyl, imidazolemethyl, pyridylmethyl and 2-pyridylethyl; wherein each of R5 and R8 is independently selected from

 $-\left(CH_{2}\right)_{q} \begin{bmatrix} R_{13} \\ C \\ R_{14} \end{bmatrix}_{p}$ Q = C-V

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wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, methyl and ethynyl; wherein R₇ is cyclohexylmethyl; wherein each of R₄ and R₆ is independently selected from hydrido and methyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl and phenyl; wherein m is

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zero; wherein n is a number selected from zero through three; wherein p is a number selected from one through three; and wherein c is zero or one; or a pharmaceutically-acceptable salt thereof.

29. The method of Claim 28 wherein said compound is selected from compounds, their tautomers, and the pharmaceutically-acceptable esters and salts thereof, of the group consisting of

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30. The method of Claim 28 wherein said compound is N1-[1R*-[[1S,1R*+(cyclohexylmethyl)-2S*,3R*-dihydroxy-hexynyl]amino]carbonyl]-3-butynyl]-N4-[2-(dimethylamino)ethyl]-N4-methyl-2S*-(phenylmethyl)butanediamide or a pharmaceutically-acceptable salt thereof.

31. The method of Claim 28 wherein said compound is [1R*-[[[1R*-[[[1S,1R*-(cyclohexylmethyl)-2S*,3R*-dihydroxy-hexynyl]amino]carbonyl]-3-butynyl]amino]carbonyl]-2-phenylethyl)[2-(dimethylamino)ethyl]methylcarbamate or a pharmaceutically-acceptable salt thereof.

32. The method of Claim 28 wherein said compound is

33. The method of Claim 28 wherein said compound is

or a pharmaceutically-acceptable salt thereof.

- 34. The method of Claim 23 wherein said circulatory disorder is a cardiovascular disorder.
- 35. The method of Claim 34 wherein said cardiovascular disorder is hypertension.
 - 36. The method of Claim 23 wherein said circulatory-related disorder is glaucoma

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